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=> d 11

L1 HAS NO ANSWERS

L1 STR

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=>

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L2 STRUCTURE UPLOADED

=> s 11

SAMPLE SEARCH INITIATED 08:27:11 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 1057 TO ITERATE

100.0% PROCESSED 1057 ITERATIONS 2 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE** BATCH **COMPLETE**

PROJECTED ITERATIONS: 19190 TO 23090 2 TO 124 PROJECTED ANSWERS:

L3 2 SEA SSS SAM L1

=> search 12

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80.1% PROCESSED 2000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

5 ANSWERS

78 ANSWERS

PROJECTED ITERATIONS: 46943 TO 52937 PROJECTED ANSWERS: 5 TO 273

5 SEA SSS SAM L2

=> search 11

ENTER TYPE OF SEARCH (SSS), CSS, FAMILY, OR EXACT:. ENTER SCOPE OF SEARCH (SAMPLE), FULL, RANGE, OR SUBSET: full FULL SEARCH INITIATED 08:27:43 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 21651 TO ITERATE

100.0% PROCESSED 21651 ITERATIONS

SEARCH TIME: 00.00.01

L578 SEA SSS FUL L1

=> search 12

ENTER TYPE OF SEARCH (SSS), CSS, FAMILY, OR EXACT:. ENTER SCOPE OF SEARCH (SAMPLE), FULL, RANGE, OR SUBSET:full FULL SEARCH INITIATED 08:27:52 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 49956 TO ITERATE

100.0% PROCESSED 49956 ITERATIONS

108 ANSWERS

SEARCH TIME: 00.00.01

L6 108 SEA SSS FUL L2

=> s 16 not 15

30 L6 NOT L5 L7

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=> s 130

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=> s 17 L8 10 L7

=> d 18 fbib ab hitstr 1-10

- L8 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN
- AN 2003:16686 CAPLUS
- DN 138:205004
- TI Substituted Pyrazolopyridopyridazines as Orally Bioavailable Potent and Selective PDE5 Inhibitors: Potential Agents for Treatment of Erectile Dysfunction
- AU Yu, Guixue; Mason, Helen; Wu, Ximao; Wang, Jian; Chong, Saeho; Beyer, Bruce; Henwood, Andrew; Pongrac, Ronald; Seliger, Laurie; He, Bin; Normandin, Diane; Ferrer, Pam; Zhang, Rongan; Adam, Leonard; Humphrey, William G.; Krupinski, John; Macor, John E.
- CS Discovery Chemistry, Drug Metabolism and Pharmacokinetics, Princeton, NJ, 08543-5400, USA
- SO Journal of Medicinal Chemistry (2003), 46(4), 457-460 CODEN: JMCMAR; ISSN: 0022-2623
- PB American Chemical Society
- DT Journal
- LA English

OS CASREACT 138:205004

AB Novel pyrazolopyridopyridazines, e.g. I, have been prepared as potent and selective PDE5 inhibitors. I has been identified as a more potent and selective PDE5 inhibitor than sildenafil. It is as efficacious as sildenafil in in vitro and in vivo PDE5 inhibition models, and it is orally bioavailable in rats and dogs. The superior isoenzyme selectivity of I is expected to exert less adverse effects in humans when used for erectile dysfunction treatment.

IT 296248-82-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation of substituted pyrazolopyridopyridazines as orally bioavailable selective PDE5 inhibitors for treatment of erectile dysfunction)

RN 296248-82-3 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-(4-pyridinylmethyl)- (CA INDEX NAME)

RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2000:688225 CAPLUS

DN 133:252445

TI Preparation of fused pyridopyridazine inhibitors of cGMP phosphodiesterase

IN Yu, Guixue; Macor, John; Chung, Hyei-jha; Humora, Michael; Katipally, Kishta; Wang, Yizhe; Kim, Soojin

PA Bristol-Myers Squibb Company, USA

SO PCT Int. Appl., 137 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PAT	CENT 1	NO.			KIN	D	DATE			APPL	ICAT	ION 1	NO.		D.	ATE	
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ΡI	WO 2000056719			A1 20000928		WO 2000-US6100					20000309							
		W:	ΑE,	AL,	ΑM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CR,	CU,
			CZ,	DE,	DK,	DM,	EE,	ES,	FΙ,	GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	IL,
			IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	KΖ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,
			MD,	MG,	MK,	MN,	MW,	MX,	NO,	NΖ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,
			SK,	SL,	ТJ,	TM,	TR,	TT,	UA,	UG,	UZ,	VN,	YU,	ZA,	ZW			
		RW:	GH,	GM,	KΕ,	LS,	MW,	SD,	SL,	SZ,	TZ,	UG,	ZW,	ΑT,	BE,	CH,	CY,	DE,
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CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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                                                              19990810
CA 2368023
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                                                           P 19990810
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                                       EP 2000-916180
                     Α1
                                                              20000309
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        IE, SI, LT, LV, FI, RO
                                       US 1999-125488P
                                                           Ρ
                                                              19990322
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AU 765128
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                                       US 1999-148009P
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CN 1161341
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                                                              20000309
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                                                           P 19990810
US 6316438
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                                       US 1999-125488P
                                                           Ρ
                                                              19990322
                                       US 1999-148009P
                                                           P 19990810
MARPAT 133:252445
The title compds. [I; Y = N, CR5; Z = N, CR6 (provided that at least one
of Y and Z = N); R1, R2 = H, halo, SR7, etc.; R3 = H, alkyl, arylalkyl; R4
= H, halo, alkyl, etc.; R5, R6 = H, halo, alkyl; R7 = H, alkyl,
cycloalkyl, etc.] and their pharmaceutically acceptable salts, inhibitors
of cGMP PDE, especially type 5, useful in treating cardiovascular and sexual
disorders, were prepared E.g., a multi-step synthesis of I [Y = N; Z = CH;
R1 = 4-hydroxypiperidin-1-y1; R2 = (3-C1-4-MeOC6H3)CH2NH; R3 = Et; R4 = H]
was given. Compds. I are effective at 0.05-100 mg/kg/day.
296248-82-3P 296248-95-8P 296249-08-6P
296249-10-0P 296249-11-1P 296249-30-4P
296249-33-7P 296249-43-9P 296249-44-0P
296249-48-4P 296249-59-7P 296249-62-2P
296249-68-8P 296249-79-1P 296249-80-4P
296249-84-8P 296249-85-9P 296249-87-1P
296250-40-3P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological
study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)
   (preparation of fused pyridopyridazine inhibitors of cGMP phosphodiesterase)
296248-82-3 CAPLUS
3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine,
N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-(4-pyridinylmethyl)- (CA
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OS

AΒ

ΙT

RN

CN

INDEX NAME)

RN 296248-95-8 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N6,N9-bis[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl- (CA INDEX NAME)

RN 296249-08-6 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[(1-oxido-4-pyridinyl)methyl]- (CA INDEX NAME)

RN 296249-10-0 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-(2-pyridinylmethyl)- (CA INDEX NAME)

RN 296249-11-1 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-(3-pyridinylmethyl)- (CA INDEX NAME)

RN 296249-30-4 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[3-(4-methyl-1-piperazinyl)propyl]- (CA INDEX NAME)

MeO
$$CH_2-NH$$
 N N $NH-(CH_2)_3-N$ N

RN 296249-33-7 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[2-(4-morpholinyl)ethyl]-

(CA INDEX NAME)

RN 296249-43-9 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-N6-[(2,6-dichloro-4-pyridinyl)methyl]-3-ethyl- (CA INDEX NAME)

RN 296249-44-0 CAPLUS

CN Ethanone, 1-[4-[[[9-[[(3-chloro-4-methoxyphenyl)methyl]amino]-3-ethyl-3H-pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazin-6-yl]amino]methyl]-1-piperidinyl]- (CA INDEX NAME)

RN 296249-48-4 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[3-(1H-imidazol-1-yl)propyl]- (CA INDEX NAME)

$$\begin{array}{c|c} & \text{C1} \\ & \text{MeO} \\ \hline \\ & \text{CH}_2 - \text{NH} \\ \hline \\ & \text{N} \\ \hline \\ & \text{N} \\ \hline \\ & \text{N} \\ \hline \\ & \text{Et} \\ \end{array}$$

RN 296249-59-7 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[(tetrahydro-2-furanyl)methyl]- (CA INDEX NAME)

RN 296249-62-2 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-(4-piperidinylmethyl)-(CA INDEX NAME)

RN 296249-68-8 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[2-(4-methyl-1-piperazinyl)ethyl]- (CA INDEX NAME)

RN 296249-79-1 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[[(2R)-tetrahydro-2-furanyl]methyl]- (CA INDEX NAME)

Absolute stereochemistry.

RN 296249-80-4 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[[(2S)-tetrahydro-2-furanyl]methyl]- (CA INDEX NAME)

Absolute stereochemistry.

RN 296249-84-8 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine,
N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[3-(4-morpholinyl)propyl](CA INDEX NAME)

MeO
$$CH_2-NH$$
 N $NH-(CH_2)_3-N$ Et

RN 296249-85-9 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[2-(1-piperidinyl)ethyl]-(CA INDEX NAME)

RN 296249-87-1 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[2-(tetrahydro-2H-pyran-4-

yl)ethyl]- (CA INDEX NAME)

RN 296250-40-3 CAPLUS

CN 3H-Pyrazolo[4',3':5,6]pyrido[3,4-d]pyridazine-6,9-diamine, N9-[(3-chloro-4-methoxyphenyl)methyl]-3-ethyl-N6-[2-(1H-imidazol-5-yl)ethyl]- (CA INDEX NAME)

RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1983:612481 CAPLUS

DN 99:212481

OREF 99:32699a,32702a

TI Condensed pyridazines. I. Reaction of 5,8-dichloropyrido[2,3-d]pyridazine with carbanion

AU Oishi, Etsuo; Watanabe, Hiromi; Hayashi, Eisaku

CS Shizuoka Coll. Pharm., Shizuoka, 422, Japan

SO Yakugaku Zasshi (1983), 103(6), 623-30 CODEN: YKKZAJ; ISSN: 0031-6903

DT Journal

LA Japanese

OS CASREACT 99:212481

AB Reaction of the title compound I (R = R1 = C1) (II) with EtO2CCH2CN, MeCOCH2CO2Et, CH2(CO2Et)2, CH2(CN)2, and PhCH2CN in C6H6 in the presence of NaNH2 gave I [R = EtO2CCHCN, CH(CN)2; R1 = C1] and I [R = C1; R1 = EtO2CCHCN, CH2CO2Et, CH(CO2Et)2, PhCHCN]. II reacted with MeCOPh and EtCOPh in PhMe in the presence of NaH to give I (R = R1 = CH2COPh; R = C1,

L8 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1976:586482 CAPLUS

DN 85:186482

OREF 85:29744h,29745a

TI Structure-activity relations of the diuretic activity of triaza- and tetraazanaphthalene compounds

AU Nishikawa, Kohei; Shimakawa, Hisao; Inada, Yoshiyuki; Shibouta, Yumiko; Kikuchi, Shintaro; Yurugi, Shojiro; Oka, Yoshikazu

CS Cent. Res. Div., Takeda Chem. Ind., Ltd., Osaka, Japan

SO Chemical & Pharmaceutical Bulletin (1976), 24(9), 2057-77 CODEN: CPBTAL; ISSN: 0009-2363

DT Journal

LA English

AB The diuretic activity of 219 nitrogen containing heterocyclic compds., classified into 13 groups based on the structural features, was studied in saline loaded rats. Of the compds. studied, 104 were active at oral doses of 10-30 mg/kg. Several of the pyrimidopyridazines, pyridazinopyridazines and pyridopyridazines produced as potent diuresis and natriuresis as hydrochlorothiazide [58-93-5] at the oral dose of 0.1 mg/kg; DS 210 (I) [33222-18-3] and DS 511 (II) [39632-88-7] were selected for more extensive evaluation as diuretic agents. Structure-activity relations of the tested compds. are discussed.

IT 33222-21-8 39632-89-8 61098-88-2 RL: BIOL (Biological study)

(diuretic)

RN 33222-21-8 CAPLUS

CN Pyrimido[4,5-d]pyridazine-5,8-diamine, 2-phenyl-N5,N8-bis(phenylmethyl)-(CA INDEX NAME)

RN 39632-89-8 CAPLUS

CN Pyrido[3,4-d]pyridazine-1,4-diamine, 7-phenyl-N1,N4-bis(phenylmethyl)-(CA INDEX NAME)

RN 61098-88-2 CAPLUS

Pyrido[2,3-d]pyridazine-5,8-diamine, 2-phenyl-N5,N8-bis(phenylmethyl)-CN (CA INDEX NAME)

ANSWER 5 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN L8

1976:17256 CAPLUS ΑN

84:17256 DN

OREF 84:2855a,2858a

Syntheses of N-heterocyclic compounds. XXV. Syntheses of ΤI pyrido[3,4-d]pyridazine derivatives. 2

Oka, Yoshikazu; Omura, Kiyoshi; Miyake, Akio; Itoh, Katsumi; Tomimoto, ΑU Mitsumi; Tada, Norio; Yurugi, Shojiro Med. Res. Lab., Takeda Chem. Ind., Ltd., Osaka, Japan

CS

SO Chemical & Pharmaceutical Bulletin (1975), 23(10), 2239-50 CODEN: CPBTAL; ISSN: 0009-2363

DTJournal

LA English

OS CASREACT 84:17256 AB Twenty-nine derivs. of the potent diuretic 1,4-dimorpholino-7-phenylpyrido[3,4-d]pyridazine, e.g., I (R = Ph, Me-, Cl-, O2N-, and MeOC6H4, xylyl, 2-furyl, 2-pyridyl, 1-, 2-naphthyl; R1 = H, Me, PhCH2; R2 = morpholino, piperidino, pyrrolidino), were prepared In 1,4-dichloropyrido[3,4-d]pyridazine the 4-chloro group was more reactive toward nucleophilic substitution than the 1-chloro group. Some reaction of I, e.g. acid hydrolysis, reduction and Grignard addition reaction were also carried out. Significance of the ring N at the 6-position in I for diuretic activity is discussed.

IT 57961-44-1P

RN 57961-44-1 CAPLUS

CN Pyrido[3,4-d]pyridazine, 7-phenyl-1,4-bis[(phenylmethyl)thio]- (CA INDEX NAME)

L8 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1973:58358 CAPLUS

DN 78:58358

OREF 78:9259a,9262a

TI Syntheses of N-heterocyclic compounds. IX. Reduction of 2-aryl-5,8-disubstituted pyrimido[4,5-d]pyridazine

AU Yurugi, Shojiro; Fushimi, Tomiyoshi; Hieda, Masaru

CS Res. Dev. Div., Takeda Chem. Ind., Ltd., Osaka, Japan

SO Yakugaku Zasshi (1972), 92(11), 1316-21 CODEN: YKKZAJ; ISSN: 0031-6903

DT Journal

LA Japanese

AB Reduction of 2-aryl-5,8-disubstituted pyrimido[4,5-d]pyridazine ((I) (R = iso-PrNH, PhNH, PhCH2NH, piperidino, morpholino, etc.; R1 = Ph, m-tolyl, p-ClC6H4, 2-thienyl, 5-morpholino-2-furyl)) to 2-aryl-3,4-dihydro-5,8-disubstituted-pyrimido[4,5-d]pyridazine (II) was carried out with NaBH4, LiAlH4, sodium isopentoxide, and a catalyst. Acylation of II gave 2-aryl-3-acyl-3,4-dihydro-5,8-dimorpholinopyrimido[4,5-d]pyridazine and alkylation of II gave 2-aryl-3-alkyl-3,4-dihydro-5,8-dimorpholinopyrimido[4,5-d]pyridazine. 2-Aryl-5,8-disubstituted-3,4-dihydropyrimido-[4,5-d]pyridazines showed a strong diuretic activity.

IT 33222-21-8

RN 33222-21-8 CAPLUS

CN Pyrimido[4,5-d]pyridazine-5,8-diamine, 2-phenyl-N5,N8-bis(phenylmethyl)-(CA INDEX NAME)

ANSWER 7 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN L8

ΑN 1973:43400 CAPLUS

DN 78:43400

OREF 78:6863a,6866a

Syntheses of N-heterocyclic compounds. XII. Syntheses of ΤI pyrido[3,4-d]pyridazine and pyrido[2,3-d]pyridazine derivatives

ΑU Yurugi, Shojiro; Fushimi, Tomiyoshi; Sugihara, Hirosada; Hieda, Masaru

Res. Dev. Div., Takeda Chem. Ind., Ltd., Osaka, Japan Yakugaku Zasshi (1972), 92(11), 1333-8 CS

SO CODEN: YKKZAJ; ISSN: 0031-6903

DT Journal

LA Japanese

AΒ 1,2,3,4-Tetrahydro-7-phenylpyrido[3,4-d]pyridazine-1,4-dione (I) and 2-phenyl-5,6,7,8-tetrahydropyrido[2,3-d]pyridazine-5,8-dione (II) were converted to the corresponding dichlorides, which reacted with amines to give 1,4-bis(substituted amino)-7-phenylpyrido[3,4-d]pyridazines and 2-phenyl-5,8-bis-(substituted amino)pyrido[2,3-d]pyridazines. 1,4-Dimorpholino-7-phenylpyrido[3,4-d]pyridazine and 2-phenyl-5, 8-bis(isopropylamino)pyrido[2,3-d]pyridazine were diuretics.

ΙT 39632-89-8P 39632-90-1P RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)

RN 39632-89-8 CAPLUS

CN Pyrido[3,4-d]pyridazine-1,4-diamine, 7-phenyl-N1,N4-bis(phenylmethyl)-(CA INDEX NAME)

39632-90-1 CAPLUS RN

CN Pyrido[2,3-d]pyridazine-5,8-diamine, 2-phenyl-N5,N8-bis(phenylmethyl)-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

L8 ANSWER 8 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1972:514340 CAPLUS

DN 77:114340

OREF 77:18841a,18844a

TI Synthesis of N-heterocyclic compounds. VII. 2-Aryl-5,8-disubstituted pyrimido[4,5-d]pyridazine

AU Yurugi, Shojiro; Hieda, Masaru; Fushimi, Tomiyoshi; Kawamatsu, Yutaka; Sugihara, Hirosada; Tomimoto, Mitsumi

CS Res. Dev. Div., Takeda Chem. Ind., Ltd., Osaka, Japan

SO Chemical & Pharmaceutical Bulletin (1972), 20(7), 1528-35 CODEN: CPBTAL; ISSN: 0009-2363

DT Journal

LA English

When 2-aryl-5,8-dichloropyrimido[4,5-d]pyridazine (I, R = R1 = C1) was reacted with nucleophiles, such as amines, sodium methoxide, sodium azide, sodium sulfide etc., 5,8-disubstituted I (R = R1 = PrNH, MeS, morpholino, etc. Ar = Ph, MeC6H4, 2-pyridyl, etc.) were obtained. treatment of 2-phenyl-5-chloro-8-morpholinopyrimido-[4,5-d]pyridazine or 2-phenyl-5-morpholino-8-chloropyrimido-[4,5-d]pyridazine with nucleophiles gave I (R = R1). The phenyl group at the 2-position accelerated the substitution at position 5 and 8. The reaction of 2-phenyl-5,8-bis(substituted thio)pyrimido[4,5-d]pyridazine with Cl gave I (R = R1 = C1, Ar = Ph). Several compds. showed diuretic activity.

RN 33222-21-8 CAPLUS

CN Pyrimido[4,5-d]pyridazine-5,8-diamine, 2-phenyl-N5,N8-bis(phenylmethyl)-(CA INDEX NAME)

38277-18-8 CAPLUS RN

Pyrimido[4,5-d]pyridazine, 2-phenyl-5,8-bis[(phenylmethyl)thio]- (CA CN INDEX NAME)

ANSWER 9 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN L8

ΑN 1971:476832 CAPLUS

DN 75:76832

OREF 75:12171a,12174a

ΤI Pyrimido[4,5-d]pyridazine derivatives

Yurugi, Shojiro; Kikuchi, Shintaro Takeda Chemical Industries, Ltd. IN

PA

SO Ger. Offen., 33 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 1

FAN	FAN.CNT 1 PATENT NO.		DATE	APPLICATION NO.	DATE	
ΡI	DE 2046577	 А	19710506	DE 1970-2046577		19700922
				JP 1969-76125	Α	19690924
				JP 1970-54984	Α	19700624
	JP 48011116	В	19730410	JP 1970-54984		19700624
	NL 7014065	A	19710326	NL 1970-14065		19700923
				JP 1969-76125	Α	19690924
				JP 1970-54984	Α	19700624
	FR 2070085	A5	19710910	FR 1970-34490		19700923
	FR 2070085	B1	19750606			
				JP 1969-76125	Α	19690924
	SE 363831	В	19740204	SE 1970-12956		19700923
				JP 1969-76125	Α	19690924
				JP 1970-54984	Α	19700624
	HU 164522	В	19740228	HU 1970-TA1090		19700923
				JP 1969-76125	Α	19690924
				JP 1970-54984	Α	19700624
	NO 129907	В	19740610	NO 1970-3617		19700923
				JP 1969-76125	Α	19690924
				JP 1970-54984	Α	
	PL 81368	B1	19750830	PL 1970-143378		19700923
				JP 1969-76125	Α	19690924
				JP 1970-54984	А	19700624
	AT 299218	В	19720612	AT 1970-8633		19700924
				JP 1969-76125	Α	
				JP 1970-54984	Α	19700624
	DK 125472	В	19730226	DK 1970-4879		19700924
				JP 1969-76125	А	19690924

			JP 1970-54984	A	19700624
GB 1325769	A	19730808	GB 1970-45525		19700924
			JP 1969-76125	A	19690924
			JP 1970-54984	A	19700624
GB 1325770	A	19730808	GB 1973-11962		19700924
			JP 1969-76125	A	19690924
			JP 1970-54984	A	19700624
US 3764598	A	19731009	US 1970-75294		19700924
			JP 1969-76125	A	19690924
			JP 1970-54984	А	19700624
CH 547298	A	19740329	CH 1970-14165		19700924
			JP 1969-76125	А	19690924
			JP 1970-54984	A	19700624
CA 955941	A1	19741008	CA 1970-94012		19700924
			JP 1969-76125	А	19690924
			JP 1970-54984	А	19700624
FR 2108518	A5	19720519	FR 1971-34438		19710924
			AT 1970-8633	A	19700924

AB The diuretic title compds. (I) were prepared in a five-step reaction. Thus, benzamidine hydrochloride and [(ethylethoxy)methylene]oxalacetate were condensed in MeOH with MeONa to give 2-phenyl-4,5-bis(ethoxycarbonyl)pyrimidine, which was refluxed with NH2NH2.-H2O in MeOH to give I (R1 = Ph, R2 = ONH2NH3, R3 = OH) (II). II in aqueous HCl was stirred at room temperature to give I (R1 = Ph, R2 = R3 = OH)

(III). A mixture of III, POC13, and PC15 was heated 3 hr to give I (R1 = Ph, R2 = R3 = C1), was heated with morpholine for 3 hr at $80-5^{\circ}$ to give I (R1 = Ph, R2 = R3 = morpholino). Similarly prepared were .apprx.15 more I (R2 = R3 = aminosubstituted) and their corresponding intermediates. IT 33222-21-8P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)

RN 33222-21-8 CAPLUS

CN Pyrimido[4,5-d]pyridazine-5,8-diamine, 2-phenyl-N5,N8-bis(phenylmethyl)-(CA INDEX NAME)

L8 ANSWER 10 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1967:37890 CAPLUS

DN 66:37890

OREF 66:7227a,7230a

TI The synthesis of pyrazino[2,3-d]pyridazine and some of its derivatives

AU Patel, Natubhai R.; Castle, Raymond N.

CS Univ. of New Mexico, Albuquerque, NM, USA

SO Journal of Heterocyclic Chemistry (1966), 3(4), 512-17 CODEN: JHTCAD; ISSN: 0022-152X

DT Journal

LA English

Pyrazino[2,3-d]pyridazine (I) was synthesized by two different routes.

5,8-Dihydroxypyrazino-[2,3-d]pyridazine was converted to

5,8-Dihydroxypyrazino-[2,3-d]pyridazine (II) and

5,8-dibromopyrazino[2,3-d]pyridazine. When II was treated with various benzyl mercaptans and alk-oxides the corresponding disubstituted drivs. were obtained. II when allowed to react with aromatic amines gave 5,8-diaminopyrazino[2,3-d]pyridazines; however, with aliphatic amines only mono-substituted products were obtained substituted in the 8-position. The reaction of pyrazine-2,3-dinitrile with hydrazine gave 5,8-diaminopyrazino[2,3-d]pyridazine.

IT 13480-47-2P 13480-48-3P 13480-49-4P

RL: SPN (Synthetic preparation): PREP (Preparation)

RN 13480-47-2 CAPLUS

CN Pyrazino[2,3-d]pyridazine, 5,8-bis[(phenylmethyl)thio]- (CA INDEX NAME)

RN 13480-48-3 CAPLUS

CN Pyrazino[2,3-d]pyridazine, 5,8-bis[[(4-chlorophenyl)methyl]thio]- (CA INDEX NAME)

PAGE 2-A

RN 13480-49-4 CAPLUS

CN Pyrazino[2,3-d]pyridazine, 5,8-bis[[(3,4-dichlorophenyl)methyl]thio]- (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

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